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## SEARCH REQUEST FORM

Scien	itific and Technical	Information Center
Requester's Full Name:  Art Unit:  Mail Box and Bldg/Room Location  Mill (M)  If more than one search is submitted.	mber 30 <u>X - 11, 3 1</u> 107, C M Resuled, please prioritize	*************
Include the elected species or structures, key	words, synonyms, acrony at may have a special mea	s specifically as possible the subject matter to be searched.  ons, and registry numbers, and combine with the concept or aning. Give examples or relevant citations, authors, etc, if abstract.
Titles of Invention:	OLOUARE,	MILAS Pressure w/(-)-11/4 daxycolombe o.
Earliest Priority Filing Date: 3	FEB 200	
*For Sequence Searches Only* Please include appropriate serial number.	all pertinent information (p	parent, child, divisional, or issued patent numbers) along with the
Meare	plarely	Jaims 2,3 and 6
		Point of Contact: Barb O'Bryen Technical Information Specialist STIC CM1 6A05 308-4291
		· Talk in the same of the same
		Account.
******************	****	**********
STAFF USE ONLY	Type of Search  NA Sequence (#)	Vendors and cost where applicable
Searcher Phone #:	AA Sequence (#)	Dialog
Searcher Location:	Structure (#)	Questel/Orbit
Date Searcher Picked Up:	Bibliographic	Dr.Link
Date Completed: 10-2-02	Litigation	Lexis/Nexis
Searcher Prep & Review Time:	Fulltext	Sequence Systems
Clerical Prep Time:	Patent Family	Other (specify)

PTO-1590 (1-2000)

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=> fil reg; d stat que 14

FILE 'REGISTRY' ENTERED AT 12:40:15 ON 02 OCT 2002

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

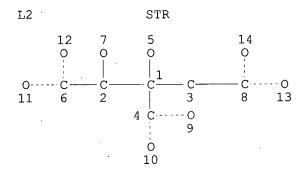
STRUCTURE FILE UPDATES: 1 OCT 2002 HIGHEST RN 457857-22-6 DICTIONARY FILE UPDATES: 1 OCT 2002 HIGHEST RN 457857-22-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf



family search done on structure of (-) hydroxy citric acid to pick up salts, stereoisomers, isotopically habelled forms & multicomponent substances

NODE ATTRIBUTES: DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

L4 34 SEA FILE=REGISTRY FAM FUL L2

100.0% PROCESSED 272 ITERATIONS

SEARCH TIME: 00.00.02

34 ANSWERS

=> fil capl; d que nos 126; d que nos 131; d que nos 134

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Page 2

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FILE COVERS 1907 - 2 Oct 2002 VOL 137 ISS 14 FILE LAST UPDATED: 1 Oct 2002 (20021001/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

```
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L6
            177 SEA FILE=CAPLUS ABB=ON L4
\Gamma8
          80941 SEA FILE=CAPLUS ABB=ON
                                         ?HYPERTENSI?
L10
          31765 SEA FILE=CAPLUS ABB=ON
                                         GLUCOCORTICOID# OR GLUCOCORTICOSTEROID#
                 OR GLYCOCORTICOID# OR CORTICOSTEROID#(L)GLUCO
T.17
          27234 SEA FILE=CAPLUS ABB=ON
                                         BLOOD PRESSURE/CT
L26
              4 SEA FILE=CAPLUS ABB=ON L6 AND (L8 OR L10 OR L17)
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L5
              1 SEA FILE=REGISTRY ABB=ON INSULIN/CN
L6
            177 SEA FILE=CAPLUS ABB=ON
                                         L4
L11
          76446 SEA FILE=CAPLUS ABB=ON
L12
         144221 SEA FILE=CAPLUS ABB=ON
                                         INSULIN
L30
          30970 SEA FILE=CAPLUS ABB=ON
                                         (L11 OR L12) (L) (RELEAS? OR SECRET? OR
                LOWER? OR DECREAS? OR INHIBIT? OR REDUC? OR PROD? OR LEVEL#)/OB
L31
              2 SEA FILE=CAPLUS ABB=ON L6 AND L30
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L5
              1 SEA FILE=REGISTRY ABB=ON INSULIN/CN
L6
            177 SEA FILE=CAPLUS ABB=ON
                                        L4
L11
          76446 SEA FILE=CAPLUS ABB=ON
L12
         144221 SEA FILE=CAPLUS ABB=ON
                                         INSULIN
L33
          10988 SEA FILE=CAPLUS ABB=ON
                                         L11(L) POTENTIAT? OR L12(W) RESISTAN?
L34
              3 SEA FILE=CAPLUS ABB=ON
                                        L33 AND L6
=> s 126 or 131 or 134
```

L73 9 L26 OR L31 OR L34

=> fil uspatf; d que nos 144

FILE 'USPATFULL' ENTERED AT 12:40:18 ON 02 OCT 2002 CA INDEXING COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS) FILE COVERS 1971 TO PATENT PUBLICATION DATE: 1 Oct 2002 (20021001/PD)
FILE LAST UPDATED: 1 Oct 2002 (20021001/ED)
HIGHEST GRANTED PATENT NUMBER: US6460183
HIGHEST APPLICATION PUBLICATION NUMBER: US2002138890
CA INDEXING IS CURRENT THROUGH 1 Oct 2002 (20021001/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 1 Oct 2002 (20021001/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2002
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2002

USPAT2 is now available. USPATFULL contains full text of the <<< >>> original, i.e., the earliest published granted patents or >>> <<< applications. USPAT2 contains full text of the latest US <<< >>> publications, starting in 2001, for the inventions covered in >>> <<< USPATFULL. A USPATFULL record contains not only the original >>> <<< published document but also a list of any subsequent >>> <<< publications. The publication number, patent kind code, and <<< >>> publication date for all the US publications for an invention >>> <<< are displayed in the PI (Patent Information) field of USPATFULL <<< >>> records and may be searched in standard search fields, e.g., /PN, <<< >>> /PK, etc. <<< >>> >>> USPATFULL and USPAT2 can be accessed and searched together <<< through the new cluster USPATALL. Type FILE USPATALL to >>> <<< >>> enter this cluster. <<< >>> <<< >>> Use USPATALL when searching terms such as patent assignees, <<< >>> classifications, or claims, that may potentially change from <<< the earliest to the latest publication. <<<

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
L2
                STR
             34 SEA FILE=REGISTRY FAM FUL L2
L4
L35
             27 SEA FILE=USPATFULL ABB=ON L4
            780 SEA FILE=USPATFULL ABB=ON
                                            (GLUCOCORTICOID# OR GLUCOCORTICOSTER
L38
                OID# OR GLYCOCORTICOID# OR CORTICOSTEROID#(L)GLUCO)/TI,IT,AB,CL
L39
           8402 SEA FILE=USPATFULL ABB=ON
                                            (ANTIHYPERTENS? OR HYPERTENS?)/TI, IT
                , AB, CLM
L40
           3647 SEA FILE=USPATFULL ABB=ON
                                            (BLOOD PRESSURE)/TI, IT, AB, CLM
L41
           5353 SEA FILE=USPATFULL ABB=ON
                                            INSULIN/TI, IT, AB, CLM
L42
           1686 SEA FILE-USPATFULL ABB-ON L41(5A) (RELEAS? OR SECRET? OR
                LOWER? OR DECREAS? OR INHIBIT? OR REDUC? OR PROD? OR LEVEL# OR
                RESISTAN? OR POTENTIAT?)/IT, TI, AB, CLM
L44
              6 SEA FILE=USPATFULL ABB=ON L35 AND ((L38 OR L39 OR L40) OR
                L42)
```

=> fil medl; d que nos 151; d que nos 153; d que nos 168

FILE 'MEDLINE' ENTERED AT 12:40:19 ON 02 OCT 2002

FILE LAST UPDATED: 1 OCT 2002 (20021001/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2002 vocabulary. Enter HELP THESAURUS for details.

THIS FILE CONTAINS CAS REGISTRY NUMBERS FOR EASY AND ACCURATE

#### SUBSTANCE IDENTIFICATION.

```
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L45
             70 SEA FILE=MEDLINE ABB=ON L4
L46
         154117 SEA FILE=MEDLINE ABB=ON HYPERTENSION+NT/CT
T.48
           4214 SEA FILE-MEDLINE ABB-ON HYPERINSULINEMIA/CT OR HYPERINSULINISM
                /CT
L49
         138702 SEA FILE=MEDLINE ABB=ON GLUCOCORTICOIDS+ALL/CT
L51
              2 SEA FILE=MEDLINE ABB=ON L45 AND (L46 OR L48 OR L49)
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L45
             70 SEA FILE=MEDLINE ABB=ON L4
L47
          98218 SEA FILE=MEDLINE ABB=ON
                                         INSULIN/CT
L53
              4 SEA FILE=MEDLINE ABB=ON L47(L)(SE OR BL)/CT AND L45
                                                       Subheading SE = secretion
BL : blood
1.2
                STR
T.4
             34 SEA FILE=REGISTRY FAM FUL L2
T.45
             70 SEA FILE=MEDLINE ABB=ON L4
          25566 SEA FILE=MEDLINE ABB=ON ANTIHYPERTENSIVE AGENTS/CT
1.66
L67
         166529 SEA FILE=MEDLINE ABB=ON BLOOD PRESSURE+NT/CT
1.68
            0 SEA FILE=MEDLINE ABB=ON (L66 OR L67) AND L45
```

=> s 151 or 153

L74 6 L51 OR L53

=> fil embase; d que nos 172

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FILE COVERS 1974 TO 26 Sep 2002 (20020926/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
L2
                STR
L4
             34 SEA FILE=REGISTRY FAM FUL L2
L54
            114 SEA FILE=EMBASE ABB=ON L4
         160157 SEA FILE=EMBASE ABB=ON HYPERTENSION+NT/CT
L55
L56
         216748 SEA FILE=EMBASE ABB=ON GLUCOCORTICOID+NT/CT
L57
         10484 SEA FILE=EMBASE ABB=ON INSULIN BLOOD LEVEL/CT
L69
         17887 SEA FILE=EMBASE ABB=ON ANTIHYPERTENSIVE AGENT/CT
L70
           3163 SEA FILE=EMBASE ABB=ON ANTIHYPERTENSIVE ACTIVITY/CT
L71
         132735 SEA FILE=EMBASE ABB=ON BLOOD PRESSURE+NT/CT
L72
              5 SEA FILE=EMBASE ABB=ON L54 AND ((L55 OR L56 OR L57) OR (L69
                OR L70 OR L71))
```

=> fil biosis; d que nos 165

FILE 'BIOSIS' ENTERED AT 12:40:22 ON 02 OCT 2002

Jones 09/781491 Page 5

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FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT

FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 25 September 2002 (20020925/ED)

```
L2
               STR
L4
            34 SEA FILE=REGISTRY FAM FUL L2
            27 SEA FILE=BIOSIS ABB=ON L4
L60
L61
        197253 SEA FILE=BIOSIS ABB=ON ?HYPERTENSI?
L62
        119274 SEA FILE=BIOSIS ABB=ON BLOOD PRESSURE
         33760 SEA FILE=BIOSIS ABB=ON (GLUCOCORTICOID# OR GLUCOCORTICOSTEROID
L63
               # OR GLYCOCORTICOID# OR CORTICOSTEROID#(L)GLUCO)
         196959 SEA FILE=BIOSIS ABB=ON INSULIN
L64
             2 SEA FILE=BIOSIS ABB=ON L60 AND (L61 OR L62 OR L63 OR L64)
L65
```

=> dup rem 174,173,165,172,144 FILE 'MEDLINE' ENTERED AT 12:41:31 ON 02 OCT 2002

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PROCESSING COMPLETED FOR L73
PROCESSING COMPLETED FOR L65
PROCESSING COMPLETED FOR L72
PROCESSING COMPLETED FOR L44

L75 22 DUP REM L74 L73 L65 L72 L44 (6 DUPLICATES REMOVED)

ANSWERS '1-6' FROM FILE MEDLINE ANSWERS '7-14' FROM FILE CAPLUS ANSWERS '15-16' FROM FILE BIOSIS ANSWERS '17-20' FROM FILE EMBASE ANSWERS '21-22' FROM FILE USPATFULL

=> d ibib abs hitstr 7-14; d ibib abs hitstr 21-22; d iall 1-6; d iall 15-20

L75 ANSWER 7 OF 22 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 1

ACCESSION NUMBER:

2002:655096 CAPLUS

DOCUMENT NUMBER:

137:179927

TITLE:

SOURCE:

(-)-Hydroxycitric acid for the prevention of

osteoporosis

INVENTOR(S):

Clouatre, Dallas L.; Dunn, James M.

PATENT ASSIGNEE(S):

USA

U.S., 8 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

Searched by Barb O'Bryen, STIC 308-4291

#### PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_ \_\_\_\_\_\_ -----\_\_\_\_\_ US 6441041 20020827 В1 US 2001-886499 20010620 AΒ (-)-Hydroxycitrate (HCA) supplementation constitutes a means of reducing the loss in bone mineral content such as that usually found in osteoporosis and the related loss in bone quality (protection against the corticoid-induced loss in non-mineral bone components). Similarly, HCA supplementation constitutes a means of reducing stress-induced bone loss and of reducing other forms of bone loss induced by glucocorticoid -related mechanisms. The discovery that HCA has bone loss-moderating effects allows for the creation of novel and more efficacious approaches to preventing estemporosis and for maintaining normal bone metabolic functioning even in the face of diet and exercise habits which are less than ideal and in the face of chronic stress. Furthermore, the discovery makes possible the development of adjuvant modalities which can be used to improve the results realized through the employment of conventional anti-osteoporosis/bone protective remedies. Controlled-release can be used to provide a sustained and modulated amt. of the active to the body as desired and therefore to regulate the use of the compd.

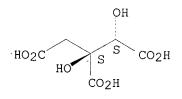
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

((-)-hydroxycitric acid for prevention of osteoporosis)

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).





RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 52729-47-2 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, trisodium salt (9CI) (CA INDEX NAME)

●3 Na

RN 64913-19-5 CAPLUS

CN Pentaric acid, 3-C-carboxy-2-deoxy-, sodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

●x Na

RN 132436-67-0 CAPLUS

CN D-threo-Pentaric acid, 3-C-carboxy-2-deoxy-, magnesium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

●x Mg

RN 185196-38-7 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, potassium salt (9CI) (CA INDEX NAME)

●x K

RN 213385-58-1 CAPLUS

Absolute stereochemistry. Rotation (-)

●x Ca

RN 449158-84-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, calcium potassium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

•x Ca

•x K

REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L75 ANSWER 8 OF 22 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 2 ACCESSION NUMBER: 2002:425329 CAPLUS

DOCUMENT NUMBER: 136:40688

TITLE: Compositions and methods for regulating metabolism and

balancing body weight

INVENTOR(S): Yegorova, Inna; Jiang, David

Searched by Barb O'Bryen, STIC 308-4291

PATENT ASSIGNEE(S):

Braswell, A. Glenn, USA

SOURCE:

U.S., 9 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_\_ \_\_\_\_\_ \_\_\_\_\_ US 6399089 20020604 US 2000-571327 В1 20000515

Compns. and methods for balancing body-wt. by inhibiting re-uptake of AΒ serotonin, regulating metab., potentiating insulia, and inhibiting lipogenesis, in a mammal. The comprise chromium, fat-free cocoa powder, Hypericum perforatum ext., Garcinia cambogia ext., Ginkgo biloba ext., Panax ginseng ext., and quercetin. For example, compn. contg. 100 .mu-g chromium, 125 mg fat-free cocoa powder, 10 mg H. perforatum ext., 125 mg G. cambogia ext., 60 mg G. biloba ext., 40 mg P. ginseng ext., and 25 mg quercetin was prepd. in tablet form. The recommended dosage for an av. wt. adult human (70-kg) is three tablets per day. In a clin. study was conducted using prepd. tablets in men having a body mass index of > 30. An increase in phys. activity and insulin sensitivity, and a decrease in dietary intake and body mass index are obsd. in the treated subjects upon completion of the study, but not in the control subjects.

27750-10-3, Hydroxycitric acid ΤТ

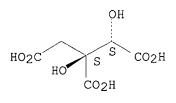
> RL: NPO (Natural product occurrence); BIOL (Biological study); OCCU (Occurrence)

(of Garcinia cambogia ext.; oral compns. contg. chromium, fat-free cocoa powder, plant exts., and quercetin for regulating/metab. and balancing body wt.)

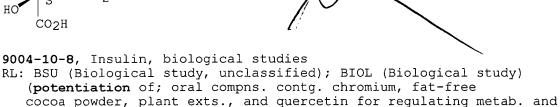
RN27750-10-3 CAPLUS

D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME) CN

Rotation (-) Absolute stereochemistry.



IΤ



balancing body wt.) 9004-10-8 CAPLUS RN

Insulin (9CI) CN (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 2 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L75 ANSWER 9 OF 22 CAPLUS COPYRIGHT 2002 ACS

2001:851796 CAPLUS

DUPLICATE 3

ACCESSION NUMBER:

135:366751

DOCUMENT NUMBER: TITLE:

Methods and pharmacoutical preparations for

normalizing blood pressure with (-)-hydroxycitric acid

INVENTOR(S): Clouatre, Dallas L.; Dunn, James M.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 6 pp. CODEN: USXXCO

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

US 2001044469 A1 20011122 US 2001-781491 20010213

PRIORITY APPLN. INFO.: US 2000-181285P P 20000209

A method whereby the blood pressure metab. in an individual showing evidence of dysregulation is improved when that person receives an appropriate oral administration of (-)-hydroxycitric acid (I). The potassium salt of I is a prefetred form of the compd., followed by the sodium salt, then by the amide and other derivs. of the acid. The regulation of blood pressure levels over any given period of time may be improved with a controlled release form of I. Controlled release can be used to provide a sustained and modulated amt. of the active to the body as desired and therefore regulate the use of the compd. as a hypotensive agent. Oral administration of 3-4 g of potassium salt of I per day in two divided doses in extremely obese patients normalized the blood pressure along with decrease of blood glucose level

IT 27750-10-3, (-)-Hydroxycitric acid 27750-10-3D,

(-)-Hydroxycitric acid, alk. earth metal salts 64913-19-5

132436-67-0 185196-38-7 213885-58-1

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Riological study); USES (Uses)

(methods and pharmaceutical prepns. for normalizing blood pressure with hydroxycitric acid salts)

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 64913-19-5 CAPLUS

CN Pentaric acid, 3-C-carboxy-2-deoxy-, sodium salt (9CI) (CA INDEX NAME)

●x Na

RN 132436-67-0 CAPLUS

Absolute stereochemistry. Rotation (-).

●x Mg

RN 185196-38-7 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, potassium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

•x K

RN 213385-58-1 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, calcium salt (9CI) (CA INDEX NAME)

●x Ca

L75 ANSWER 10 OF 22 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 4

ACCESSION NUMBER: 2001:224396 CAPLUS

DOCUMENT NUMBER: 134:256874

TITLE: Methods and pharmaceutical preparations for <u>impro</u>ving

glucose metabolism with (-)-hydroxycitric acid

INVENTOR(S): Clouatre, Dallas L.; Dunn, James M.

PATENT ASSIGNEE(S): USA

SOURCE: U.S., 4 pp.
CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ---------\_\_\_\_ \_\_\_\_\_ -----US 6207714 В1 20010327 US 2000-661588 20000914 PRIORITY APPLN. INFO.: US 1999-153840P P 19990914 Disclosed is a method whereby the glucose metab. in an individual showing

evidence of dysregulation, as is found in insulin resistance, reactive hyperglycemia and/or elevated blood sugar levels and/or diabetes, is improved when that person receives an appropriate oral administration of (-)-hydroxycitric acid. The potassium salt of (-)-hydroxycitric acid is the preferred form of the compd., followed by the sodium salt. The regulation of glucose levels over any given period of time may be improved with a controlled release form of (-)-hydroxycitric acid. Controlled release can be used to provide a sustained and modulated amt. of the active to the body as desired and therefore regulate the use of the compd. as a hypoglycemic agent.

IT 185196-38-7

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(improving glucose metab. with (-)-hydroxycitric acid and its salts)

RN 185196-38-7 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, potassium salt (9CI) (CA INDEX NAME)

●x K

IT 27750-10-3, (-)-Hydroxycitric acid 64913-19-5 132436-67-0 213385-58-1

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (improving glucose metab. with (-)-hydroxycitric acid and its salts)

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 64913-19-5 CAPLUS

CN Pentaric acid, 3-C-carboxy-2-deoxy-, sodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

●x Na

RN 132436-67-0 CAPLUS

CN D-threo-Pentaric acid, 3-C-carboxy-2-deoxy-, magnesium salt (9CI) (CA INDEX NAME)

●x Mg

RN 213385-58-1 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy-, calcium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

●x Ca

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L75 ANSWER 11 OF 22 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2000:336612 CAPLUS

DOCUMENT NUMBER:

133:119495

TITLE:

SOURCE:

IT

Toward a wholly nutritional therapy for type 2

diabetes

AUTHOR(S):

McCarty, M. F.

CORPORATE SOURCE:

Helicon Foundation, San Diego, CA, USA Medical Hypotheses (2000), 54(3), 483-487

CODEN: MEHYDY; ISSN: 0306-9877

PUBLISHER:
DOCUMENT TYPE:

Churchill Livingstone
Journal; General Review

LANGUAGE:

English

A review with 84 refs. is given. It may now be feasible to target specific supplemental nutrients to each of the key dysfunctions which conspire to maintain hyperglycemia in type 2 diabetes: bioactive chromium

of skeletal muscle insulin resistance, conjugated linoleic acid for adipocyte insulin resistance,

high-dose biotin for excessive hepatic glucose output, and coenzyme Q10 for beta cell failure. Nutritional strategies which disinhibit hepatic fatty acid oxidn. (involving hydroxycitrate, carnitine, pyruvate, and other adjuvants) may likewise prove beneficial - in the short term, by decreasing serum free fatty acids and, in the longer term, by promoting regression of visceral obesity. The nutrients and food factors recommended here appear to be safe and well tolerated, and thus may have particular utility for diabetes prevention.

27750-10-3, Hydroxycitric acid

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(toward a wholly nutritional therapy for type 2 diabetes)

Page 15

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

REFERENCE COUNT:

84 THERE ARE 84 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L75 ANSWER 12 OF 22 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1998:268507 CAPLUS

DOCUMENT NUMBER:

128:278299

TITLE:

Magnesium (-)-hydroxycitrate, method of preparation,

W 19971017

applications, and compositions, in particular

pharmaceutical, containing same

INVENTOR(S):

Shrivastava, Ravi; Lambropoulos, Patrick Shrivastava, Ravi, Fr.; Lambropoulos, Patrick

PATENT ASSIGNEE(S): PCT Int. Appl., 23 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent French

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	TENT NO.	K	IND DATE	E	APPLICATION	NO.	DATE		a	
WO	98 <u>1767</u> 1		A1 1998	30430	WO 1997-FR1	860	19971017			
	W: AU, RW: AT,	•		ES. FT.	FR, GB, GR, II	E. TT.	IU. MC.	NT	PT. S	F.
FR	2754820	•	A1 1998		FR 1996-130		19961022		, -	_
	2754820		31 1999		4005 405	•				
	9748717	_		30515	AU 1997-487	17	19971017			
	717533 937085	-		00330	EP 1997-911.	205	10071017			
D.F					GB, GR, IT, L			MC,	PT,	
	IE,									ı
JP	200150374	44 1	г2 2001	10321	JP 1998-519	029	19971017			L
KR	200005268	37 2	A 2000	00825	KR 1999-703	474	19990421			V
US	6221901	]	31 2001	10424	US 1999-284	864	19990422			-
PRIORITY	Y APPLN.	INFO.:			FR 1996-13094	Α	19961022	_		

AB The invention concerns magnesium (-)-hydroxycitrate, its method of prepn., its applications in dietetics and in therapeutics particularly in the cardiovascular field, and pharmaceutical compns. contq. it. Thus, magnesium (-)-hydroxycitrate is prepd. from reaction of an ext. of Garcinia cambogia with an aliph. alc. (e.g., EtOH) to obtain a ppt. which is treated with a tannin fixative (e.g., poly(vinylpyrfolidone)), filtered, and the remaining soln. agitated with an anion exchange resin, the supernatant is eliminated, and the product is eluted and dried. Magnesium (-)-hydroxycitrate is useful in the therapeutic treatment of cardiovascular diseases. The antioxidant and antihypertensive activities of the (-)-hydroxycitrate in rat, its antihypercholesterolemic and antiatherosclerotic activities in rabbit, and the toxicity in rat are reported. An assocn. of magnesium (-)-hydroxycitrate with Mg, Cu, Co, Zn, Ni, Se, Si, Mn, Li, or Fe, ionized or not, and at least one vitamin is claimed. Pharmaceutical formulations contg. magnesium (-)-hydroxycitrate

WO 1997-FR1860

are claimed (6 examples). Magnesium (-)-hydroxycitrate or an assocd. compd. described above are applicable to dietetic/nutritional or cosmetic products.

ΙT 132436-67-0P

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of magnesium (-)-hydroxycitrate for treatment of cardiovascular diseases)

RN 132436-67-0 CAPLUS

CND-threo-Pentaric acid, 3-C-carboxy-2-deoxy-, magnesium salt (9CI) INDEX NAME)

Absolute stereochemistry. Rotation (-).

#### x Mg

L75 ANSWER 13 OF 22 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:784225 CAPLUS

DOCUMENT NUMBER: 130:177001

TITLE: Utility of metformin as an adjunct to

hydroxycitrate/carnitine for reducing body fat in

diabetics

AUTHOR(S): McCarty, M. F.

CORPORATE SOURCE: Nutrition 21, San Diego, CA, 92109, USA SOURCE: Medical Hypotheses (1998) 51(5), 399-403

CODEN: MEHYDY; ISSN: 0306-9877

PUBLISHER: Churchill Livingstone DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

A review with 39 refs. Excessive exposure of tissues to fatty acids is likely to be the chief cause of the various dysfunctions that lead to sustained hyperglycemia in type II diabetes. These dysfunctions are likely to be substantially reversible if body fat and dietary fat can be greatly reduced. Disinhibition of Appatic fatty acid oxidn, with hydroxycitrate (HCA) and carnitine has considerable potential as a new wt.-loss strategy, but in diabetics runs the risk of further enhancing excessive hepatic gluconeogenesis. Since the clin. utility of metformin in diabetes is probably traceable to inhibition of gluconeogenesis, its use as an adjunct to HCA/carnitine treatment of obesity in diabetics deserves evaluation, particularly as metformin therapy itself tends to reduce body wt. A consideration of relevant evidence suggests that metformin therapy will not impede the activation of fatty acid oxidn. by HCA/carnitine, and is likely to potentiate the appetite-suppressant and thermogenic benefits of this strategy. Indeed, since metformin has been reported to lower body wt. and improve cardiovascular risk factors in obese non-diabetics, a broader application of a metformin/HCA/carnitine therapy for obesity can be contemplated.

TΤ 27750-10-3, Hydroxycitric acid RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

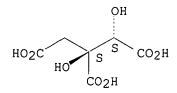
Page 17

(utility of metformin as an adjunct to hydroxycitrate/carnitine for reducing body fat in diabetics)

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 78 THERE ARE 78 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L75 ANSWER 14 OF 22 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:720056 CAPLUS

DOCUMENT NUMBER: 127:351178

TITLE: Dietary composition containing chitosan, Garcinia

cambogia hydroxycitrate, and organic chromium

INVENTOR(S):
Littera, Renato

PATENT ASSIGNEE(S): SIRC S.P.A. Natural & Dietetic Foods, Italy

SOURCE: Eur. Pat. Appl., 6 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 803202	A2	19971029	EP 1997-830189	19970424
EB 00000	7 7	10000400		

EP 803202 A3 19980429

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, SI, LT, LV, FI, RO

PRIORITY APPLN. INFO.:

AB The use of prepns. based on the combination of chitosan with org. chromium and Garcinia cambogia hydroxycitrate as dietary products for the treatment of obesity having hypocholesteremic and sugar absorption reducing activity is disclosed. The proposed combination of chitosan with org. chromium and Garcinia cambogia hydroxycitrate is formulated on the base of the effects that the above three components have on the glucid metab. Such effects tends particularly to decrease the values of cholesterolemia and triglycerides in case they are too high. The integrator of the invention can be administered by mouth in the usual dose unit both as capsules and tablets and is efficacious as diet integrator in the wt. reducing programs aiming at calorie restrictions in obese subjects, in the treatment of

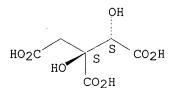
hypertension, and as hypocholesteremic product.
IT 27750-10-3, Hydroxycitric acid

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); FFD (Food or feed use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(dietary compn. contg. chitosan, Garcinia cambogia hydroxycitrate, and org. chromium)

RN 27750-10-3 CAPLUS

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)



L75 ANSWER 21 OF 22 USPATFULL

ACCESSION NUMBER:

2001:59921 USPATFULL

TITLE:

Magnesium (-) hydroxycitrate, method of preparation,

applications, and compositions in particular

pharmaceutical containing same

Shrivastava, Ravi, 43bis route de Chateaugay, 63118 INVENTOR(S):

Cebazat, France

Lambropoulos, Patrick, 35 Traverse Nicolas, 13007

Marseille, France

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 6221901	B1	20010424	
FAIENI INFORMATION.	WO 9817671	DI	19980430	
APPLICATION INFO.:	US 1999-284864		19990422	(9)
	WO 1997-FR1860		19971017	
				PCT 371 date /
			19990422	PCT 102(e) date
				/

NUMBER DATE

PRIORITY INFORMATION:

FR 1996-13094 19961022

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER: LEGAL REPRESENTATIVE: O'Sullivan, Peter Browdy and Neimark

NUMBER OF CLAIMS:

23 1

EXEMPLARY CLAIM:

LINE COUNT: 508

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Magnesium (-) hydroxycitrate, preparation process, dietary and therapeutic uses particularly in the cardiovascular field, and compositions in particular pharmaceutical containing it.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 132436-67-0P

(prepn. of magnesium (-)-hydroxycitrate for treatment of cardiovascular diseases)

RN 132436-67-0 USPATFULL

CN D-threo-Pentaric acid, 3-C-carboxy-2-deoxy-, magnesium salt (9CI) INDEX NAME)

●x Mg

L75 ANSWER 22 OF 22 USPATFULL

ACCESSION NUMBER: 1998:14823 USPATFULL

TITLE: Method of treatment for carbohydrate addiction INVENTOR(S): Bernstein, Richard K., 1160 Greacen Point Rd., Mamaroneck, NY, United States 10543

NUMBER KIND DATE

PATENT INFORMATION: US 5716976 19980210

APPLICATION INFO.: US 1996-615616 19960313 (8)

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Fay, Zohreh

LEGAL REPRESENTATIVE: Kane, Dalsimer, Sullivan, Kurucz, Levy, Eisele and

Richard

NUMBER OF CLAIMS: 27 EXEMPLARY CLAIM: 1 LINE COUNT: 713

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method is described for alleviating carbohydrate addiction by

administration of anorexients on a schedule that avoids tolerance to the

∕anorexient.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 27750-10-3, Hydroxycitric acid

(anorexient treatment of carbohydrate addiction)

RN 27750-10-3 USPATFULL

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L75 ANSWER 1 OF 22 MEDLINE DUPLICATE 6

ACCESSION NUMBER: 92144681 MEDLINE

DOCUMENT NUMBER: 92144681 PubMed ID: 1782221

TITLE: Hexose metabolism in pancreatic islets. Effect of

(-)-hydroxycitrate upon fatty acid synthesis and insulin

release in glucose-stimulated islets.

for Medline Biosis & Embase hits printed at end

AUTHOR: Sener A; Malaisse W J

CORPORATE SOURCE: Laboratory of Experimental Medicine, Brussels Free

University, Belgium.

SOURCE: BIOCHIMIE, (1991 Oct) 73 (10) 1287-90.

Journal code: 1264604. ISSN: 0300-9084.

PUB. COUNTRY: France

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199203

ENTRY DATE: Entered STN: 19920405

Last Updated on STN: 19920405 Entered Medline: 19920313

ABSTRACT:

Anaplerotic reactions leading to the de novo synthesis of fatty acids, were recently proposed to participate in the coupling of metabolic to secretory events in the process of glucose-stimulated insulin release. In an attempt to validate such a proposal, the effect of (-)-hydroxycitrate upon fatty acid synthesis and insulin release was investigated in glucose-stimulated hat pancreatic islets. The inhibitor of ATP citrate-lyase, when tested in the 1.0-2.0 mM range, failed to affect glucose-stimulated insulin release, but also failed to inhibit the incorporation of 14C-labelled acetyl residues derived from L-[U-14C]leucine into islet lipids. A partial inhibition of fatty acid labelling by 3H2O was only observed in islets incubated for 120 min in the presence of 5.0 mM (-)-hydroxycitrate and absence of CaCl2. These findings suggest that (-)-hydroxycitrate is not, under the present experimental conditions, a useful tool to abolish fatty acid synthesis in intact pancreatic islets.

CONTROLLED TERM: Check Tags: Animal; Female; In Vitro; Support, Non-U.S.

Gov't

Citrates: PD, pharmacology
Fatty Acids: BI, biosynthesis
Glucose: PD, pharmacology
\*Hexoses: ME, metabolism
Insulin: SE, secretion

Islets of Langerhans: DE, drug effects \*Islets of Langerhans: ME, metabolism Islets of Langerhans: SE, secretion

Rats

CAS REGISTRY NO.: 11061-68-0 (Insulin); 50-99-7 (Glucose); 6205-14-7

(hydroxycitric acid)

CHEMICAL NAME: 0 (Citrates); 0 (Fatty Acids); 0 (Hexoses)

L75 ANSWER 2 OF 22 MEDLINE

ACCESSION NUMBER: 2001105565 MEDLINE

DOCUMENT NUMBER: 20583412 PubMed ID: 11187927

TITLE: Dietary fat intake, supplements, and weight loss.

AUTHOR: Dyck D J

CORPORATE SOURCE: Department of Human Biology and Nutritional Sciences,

University of Guelph, ON.

SOURCE: CANADIAN JOURNAL OF APPLIED PHYSIOLOGY, (2000 Dec) 25 (6)

495-523. Ref: 159

Journal code: 9306274. ISSN: 1066-7814.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200102

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20010322

Jones 09/781491 Page 21

Entered Medline: 20010208

ABSTRACT:

Although there remains controversy regarding the role of macronutrient balance in the etiology of obesity, the consumption of high-fat diets appears to be strongly implicated in its development. Evidence that fat oxidation does not adjust rapidly to acute increases in dietary fat, as well as a decreased capacity to oxidize fat in the postprandial state in the obese, suggest that diets high in fat may lead to the accumulation of fat stores. Novel data is also presented suggesting that in rodents, high-fat diets may lead to the development of leptin resistance in skeletal muscle and subsequent accumulations of muscle triacylglycerol. Nevertheless, several current fad diets recommend drastically reduced carbohydrate intake, with a concurrent increase in fat content. Such recommendations are based on the underlying assumption that by reducing circulating insulin levels, lipolysis and lipid oxidation will be enhanced and fat storage reduced. Numerous supplements are purported to increase fat oxidation (carnitine, conjugated linoleic acid), increase metabolic rate (ephedrine, pyruvate), or inhibit hepatic lipogenesis (hydroxycitrate). All of these compounds are currently marketed in supplemental form to increase weight loss, but few have actually been shown to be effective in scientific studies. To date, there is little or no evidence supporting that carnitine (or) hydroxycitrate supplementation are of any value for weight loss in humans Supplements such as pyruvate have been shown to be effective at high dosages, but there is little mechanistic information to explain its purported effect or data to indicate its effectiveness at Yower dosages. Conjugated linoleic acid has been shown to stimulate fat utilization and decrease body fat content in mice but has not been tested in humans. The effects of ephedrine conjunction with methylxanthines and aspirin, in humans appears unequivocal includes various cardiovascular side effects. None of these compounds have been tested for their effectiveness or safety over prolonged periods of time.

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Check Tags: Animal; Human; Support, Non-U.S. Gov't;
CONTROLLED TERM:
                    Support, U.S. Gov't, Non-P.H.S.
                     Anti-Obesity Agents: AE, adverse effects
                     Anti-Obesity Agents: TU, therapeutic use
                     Aspirin: AE, adverse effects
                     Aspirin: TU, therapeutic use
                     Carnitine: TU, therapeutic use
                     Citrates: TU, therapeutic use
                    *Dietary Fats: AD, administration & dosage
                     Dietary Fats: AE, adverse effects
                    *Dietary Supplements
                     Dietary Supplements: AE, adverse effects
                     Ephedrine: TU, therapeutic use
                       Insulin: BL, blood
                     Leptin: ME, metabolism
                     Linoleic Acid: TU, therapeutic use
                     Lipids: ME, metabolism
                     Lipolysis
                     Mice
                     Muscle, Skeletal: ME, metabolism
                     Obesity: ET, etiology
                     Oxidation-Reduction
                     Pyruvates: TU, therapeutic use
                     Rats
                     Triglycerides: ME, metabolism
                    *Weight Loss
                     Xanthines: AE, adverse effects
                     Xanthines: TU, therapeutic use
CAS REGISTRY NO.:
                    11061-68-0 (Insulin); 2197-37-7 (Linoleic Acid); 28109-92-4
                    (methylxanthine); 299-42-3 (Ephedrine); 50-78-2 (Aspirin);
                    541-15-1 (Carnitine); 6205-14-7 (hydroxycitric
                    acid)
CHEMICAL NAME:
                    0 (Anti-Obesity Agents); 0 (Citrates); 0 (Dietary Fats); 0
```

Jones 09/781491 Page 22

(Leptin); 0 (Lipids); 0 (Pyruvates); 0 (Triglycerides); 0 (Xanthines)

L75 ANSWER 3 OF 22 MEDLINE

97344123 ACCESSION NUMBER: MEDLINE

DOCUMENT NUMBER: 97344123 PubMed ID: 9200650

Stimulation of islet protein kinase C translocation by TITLE:

palmitate requires metabolism of the fatty acid.

AUTHOR: Alcazar O; Qiu-yue Z; Gine E; Tamarit-Rodriguez J

CORPORATE SOURCE: Department of Biochemistry, Complutense University Medical

School, Madrid, Spain.

SOURCE: .

DIABETES, (1995 Jul) 46 (7) 1153-8. Journal code: 0 72763. ISSN: 0012-1797.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199707

ENTRY DATE: Entered STN: 19970724

> Last Updated on STN: 19970724 Entered Medline: 19970716

#### ABSTRACT:

The Secretory, metabolic, and signaling aspects of glucose/palmitate interaction on beta-cell function have been studied on rat islets. Palmitate potentiated the glucose-induced insulin response of perifused islets at suprathreshold (>3 mmol/l) sugar concentrations. This potentiating effect could be suppressed by 8-bromo-cGMP, which also blocks palmitate metabolism. Palmitate did not modify glucose utilization, but it slightly reduced glucose oxidation and concomitantly increased lactate production. The very low rate of palmitate oxidation (80-fold lower than that of 20 mmol/l glucose) might explain its lack of effect on glycolysis and hence that the glucose/fatty acid cycle is inoperative in islet cells. However, glucose determines the metabolic fate of exogenous palmitate, which is mainly diverted toward lipid synthesis at high sugar concentrations and might then generate lipid messengers for cell signaling. Palmitate did not increase glucose-induced production of inositol-1,4,5-trisphosphate, but it stimulated the translocation of protein kinase C activity from a cytosolic to a particulate fraction at 20 but not at 3 mmol/l glucose. This increased translocation was partially or completely blocked by hydroxycitrate or 8-bromo-gGMP, respectively, which are agents interfering with palmitate metabolism (inhibiting lipid synthesis). The metabolic interaction between glucose and palmitate might generate lipid messengers (diacylglycerol, phosphatidylserine) necessary for the activation of i<del>blet</del> protein kinase C, which would in turn result in a potentiation of glucose-induced insulin secretion.

CONTROLLED TERM:

Check Tags: Animal; Comparative Study; Male; Support,

Non-U.S. Gov't

8-Bromo Cyclic Adenosine Monophosphate: PD, pharmacology

Citrates: PD, pharmacology

Cyclic GMP: AA, analogs & derivatives

Cyclic GMP: PD, pharmacology Cytosol: EN, enzymology Cytosol: ME, metabolism

Dose-Response Relationship, Drug

\*Glucose: ME, metabolism Glucose: PD, pharmacology Insulin: IM, immunology \*Insulin: SE, secretion

Islets of Langerhans: DE, drug effects Islets of Langerhans: EN, enzymology \*Islets of Langerhans: PH, physiology

Lactic Acid: BI, biosynthesis Membrane Proteins: ME, metabolism Octanoic Acids: ME, metabolism

Oxidation-Reduction

\*Palmitates: ME, metabolism Palmitates: PD, pharmacology Protein Kinase C: DE, drug effects \*Protein Kinase C: ME, metabolism

Rats

Rats, Wistar

Rotenone: PD, pharmacology

Time Factors

CAS REGISTRY NO.: 11061-68-0 (Insulin); 124-07-2 (caprylic acid); 23583-48-4

(8-Bromo Cyclic Adenosine Monophosphate); 31356-94-2 (8-bromocyclic GMP); 50-21-5 (Lactic Acid); 50-99-7

(Glucose); **6205-14-7** (hydroxycitric acid); 7665-99-8 (Cyclic GMP); 83-79-4 (Rotenone)

CHEMICAL NAME: 0 (Citrates); 0 (Membrane Proteins); 0 (Octanoic Acids); 0

(Palmitates); EC 2.7.1.- (Protein Kinase C)

L75 ANSWER 4 OF 22 MEDLINE

ACCESSION NUMBER: 96130666 MEDLINE

DOCUMENT NUMBER: 96130666 PubMed ID: 8569547

TITLE: Inhibition of citrate lyase may aid aerobic endurance.

AUTHOR: McCarty M F

SOURCE: MEDICAL HYPOTHESES, (1995 Sep) 45 (3) 247-54. Ref: 77

Journal code: 7505668. ISSN: 0306-9877.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals; Space Life Sciences

ENTRY MONTH: 199603

ENTRY DATE: Entered STN: 19960315

Last Updated on STN: 19980206 Entered Medline: 19960305

### ABSTRACT:

Owing to a substantial increase in glucose uptake by working muscle, glucose homeostasis during sustained aerobic exercise requires a severalfold increase in hepatic glucose output. As exercise continues and liver glycogen declines, an increasing proportion of this elevated glucose output must be provided by gluconeogenesis. Increased gluconeogenic efficiency in trained individuals is a key adaptation promoting increased endurance, since failure of hepatic glucose output to keep pace with muscle uptake rapidly leads to hypoglycaemia and exhaustion. Pre-administration of (-)-hydroxycitrate, a potent inhibitor of citrate lyase found in fruits of the genus Garcinia, may aid endurance during post-absorptive aerobic exercise by promoting gluconeogenesis. Carnitine and bioactive chromium may potentiate this benefit. The utility of this technique may be greatest in exercise regimens designed to promote weight loss.

CONTROLLED TERM: Check Tags: Animal; Human

Aerobiosis

Carnitine: PD, pharmacology

Chromium Compounds: PD, pharmacology

\*Citrates: PD, pharmacology \*Exertion: PH, physiology

\*Gluconeogenesis: DE, drug effects

\*Glucose: ME, metabolism \*Glycogen: ME, metabolism

Glycolysis

Hormones: PH, physiology Lipids: ME, metabolism Liver: ME, metabolism

\*Multienzyme Complexes: AI, antagonists & inhibitors

Jones 09/781491 Page 24

Multienzyme Complexes: PH, physiology

Muscle, Skeletal: ME, metabolism

\*Oxo-Acid-Lyases: AI, antagonists & inhibitors

Oxo-Acid-Lyases: PH, physiology \*Physical Endurance: DE, drug effects

Rats

Weight Loss: DE, drug effects

CAS REGISTRY NO.: 50-99-7 (Glucose); 541-15-1 (Carnitine); 6205-14-7

(hydroxycitric acid); 9005-79-2 (Glycogen)
CHEMICAL NAME: 0 (Chromium Compounds); 0 (Citrates); 0 (Hormones); 0

(Lipids); 0 (Multienzyme Complexes); EC 4.1.3.

(Oxo-Acid-Lyases); EC 4.1.3.6 (citrate (pro-3S)-lyase)

L75 ANSWER 5 OF 22 MEDLINE

ACCESSION NUMBER: 94283727 MEDLINE

DOCUMENT NUMBER: 94283727 PubMed ID: 8013751

TITLE: More direct evidence for a malonyl-CoA-carnitine

palmitoyltransferase I interaction as a key event in

pancreatic beta-cell signaling.

AUTHOR: Chen S; Ogawa A; Ohneda M; Unger R H; Foster D W; McGarry J

D

CORPORATE SOURCE: Department of Internal Medicine, Gifford Laboratories,

University of Texas Southwestern Medical Center at Dallas

75235-8858.

CONTRACT NUMBER: DK-18575 (NIDDK)

DK-42582 (NIDDK)

SOURCE: DIABETES, (1994 Jul) 43 (7) 878-83.

Journal code: 0372763. ISSN: 0012-1797.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199407

ENTRY DATE: Entered STN: 19940810

Last Updated on STN: 19980206 Entered Medline: 19940725

ABSTRACT:

We sought to explore the emerging concept that malonyl-CoA generation, with concomitant suppression of mitochondrial carnitine palmitoyltransferase I (CPT I), represents an important component of glucose-stimulated insulin secretion (GSIS) by the pancreatic beta-cell (Prentki M, Vischer S, Glennon MC, Regazzi R, Deeney JT, Corkey BE: Malonyl-CoA and long-chain acyl-CoA esters as metabolic coupling factors in nutrient-induced insulin secretion. J Biol Chem 267:5802-5810, 1992). Accordingly, pancreases from fed rats were perfused with basal (3 mM) followed by high (20 mM) glucose in the absence or presence of 2 mM hydroxycitrate (HC), an inhibitor of ATP-citrate (CIT) lyase (the penultimate step in the glucose-->malonyl-CoA conversion). HC profoundly inhibited GSIS, whereas CIT had no effect. Inclusion of 0.5 mM palmitate in the perfusate significantly enhanced GSIS and completely offset the negative effect of HC. In isolated islets, HC stimulated [1-14C] palmitate oxidation in the presence of basal glucose and markedly obtunded the inhibitory effect of high glucose. Directional changes in 14C incorporation into phospholipids were opposite to those of 14CO2 production. At a concentration of 0.2 mM, 2-bromostearate, 2-bromopalmitate and etomoxir (all CPT I inhibitors) potentiated GSIS by the pancreas and inhibited palmitate oxidation in islets. However, at 0.05 mM, etomoxir did not influence insulin secretion but still caused significant suppression of fatty acid oxidation. The results provide more direct evidence for a pivotal role of malonyl-CoA suppression of CPT I, with attendant elevation of the cytosolic long-chain acyl-CoA concentration, in GSIS from the normal pancreatic beta-cell. (ABSTRACT TRUNCATED AT 250 WORDS)

CONTROLLED TERM: Check Tags: Animal; In Vitro; Male; Support, Non-U.S. Gov't; Support, U.S. Gov't, P.H.S.

ATP Citrate (pro-S)-Lyase: AI, antagonists & inhibitors Carnitine O-Palmitoyltransferase: AI, antagonists & inhibitors

\*Carnitine O-Palmitoyltransferase: ME, metabolism

Citrates: PD, pharmacology

Epoxy Compounds: PD, pharmacology

Glucose: PD, pharmacology

Hypoglycemic Agents: PD, pharmacology

Insulin: SE, secretion

Islets of Langerhans: DE, drug effects Islets of Langerhans: EN, enzymology \*Islets of Langerhans: PH, physiology

Kinetics

Malonyl Coenzyme A: ME, metabolism

Palmitates: PD, pharmacology

Palmitic Acid

Palmitic Acids: ME, metabolism

Rats

Rats, Sprague-Dawley \*Signal Transduction

Stearic Acids: PD, pharmacology

Time Factors

CAS REGISTRY NO.: 11061-68-0 (Insulin); 142-94-9 (2-bromostearic acid);

18263-25-7 (2-bromopalmitate); 50-99-7 (Glucose); 524-14-1

(Malonyl Coenzyme A); 57-10-3 (Palmitic Acid); 6205-14-7 (hydroxycitric acid); 82258-36-4

(etomoxir)

CHEMICAL NAME: 0 (Citrates); 0 (Epoxy Compounds); 0 (Hypoglycemic Agents);

> O (Palmitates); O (Palmitic Acids); O (Stearic Acids); EC 2.3.1.21 (Carnitine O-Palmitoyltransferase); EC 4.1.3.8

(ATP Citrate (pro-S)-Lyase)

L75 ANSWER 6 OF 22 MEDLINE

ACCESSION NUMBER: 90254190 MEDLINE

DOCUMENT NUMBER:

90254190 PubMed ID: 2160286

TITLE:

Glucocorticoid induction of fatty-acid synthase mediates the stimulatory effect of the hormone on choline-phosphate

cytidylyltransferase activity in fetal rat lung.

Xu Z X; Smart D A; Rooney S A AUTHOR:

CORPORATE SOURCE: Department of Pediatrics, Yale University School of

HD-10192 (NICHD)

Medicine, New Haven, CT.

CONTRACT NUMBER:

HL-43320 (NHLBI)

SOURCE: BIOCHIMICA ET BIOPHYSICA ACTA, (1990 May 1) 1044 (1) 70-6.

Journal code: 0217513. ISSN: 0006-3002.

PUB. COUNTRY:

Netherlands

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199006

ENTRY DATE:

Entered STN: 19900720

Last Updated on STN: 19980206 Entered Medline: 19900628

#### ABSTRACT:

Fetal lung fatty-acid synthase and choline-phosphate cytidylyltransferase activities are increased by glucocorticoids. There is evidence that the hormone increases synthesis of fatty-acid synthase but only increases the catalytic activity of the cytidylyltransferase. Free fatty acids and a number of phospholipids have been reported to stimulate cytidylyltransferase activity in several organs, including the lung. We have addressed the question of whether glucocorticoid induction of fatty-acid synthase mediates the stimulatory effect of the hormone on choline-phosphate cytidylyltransferase activity. Explants of 18-day fetal rat lung were cultured for 48 h with dexamethasone and inhibitors .

of de novo fatty acid biosynthesis (agaric acid and hydroxycitric acid) being included in the medium for the final 20 h. Dexamethasone increased the activities of fatty acid synthase and choline-phosphate cytidylyltransferase by 84% and 60%, respectively. Agaric acid and hydroxycitric acid completely abolished the stimulatory effect of the hormone on cytidylyltransferase but not on fatty-acid synthase. The inhibitors had no effect on cytidylyltransferase activity in control cultures. Fetal lung choline-phosphate cytidylyltransferase can be maximally stimulated by inclusion of phosphatidylglycerol in the assay mixture and under this condition, cytidylyltransferase activity in control and dexamethasone-treated cultures in the presence and absence of the inhibitors were all increased to the same level. Therefore, the inhibitors did not diminish the capacity of cytidylyltransferase to be fully activated. We suggest that the glucocorticoid induction of fatty-acid synthase in fetal lung results in increased synthesis of fatty acids which in turn, either as free acids or after incorporation into phospholipids, activate choline-phosphate cytidylyltransferase.

CONTROLLED TERM:

Check Tags: Animal; Female; Support, U.S. Gov't, P.H.S.

Cells, Cultured

Choline-Phosphate Cytidylyltransferase

Citrates: PD, pharmacology DNA: BI, biosynthesis

Dexamethasone: PD, pharmacology Enzyme Induction: DE, drug effects

\*Fatty Acid Synthetase Complex: BI, biosynthesis Fatty Acid Synthetase Complex: GE, genetics

Fetus

\*Glucocorticoids: PD, pharmacology

Kinetics

Lung: DE, drug effects \*Lung: EN, enzymology

Nucleotidyltransferases: GE, genetics \*Nucleotidyltransferases: ME, metabolism Phosphatidylglycerols: PD, pharmacology

Pregnancy

Rats

Rats, Inbred Strains

CAS REGISTRY NO.:

50-02-2 (Dexamethasone); 6205-14-7 (hydroxycitric

acid); 666-99-9 (agaric acid); 9007-49-2 (DNA)

CHEMICAL NAME:

0 (Citrates); 0 (Glucocorticoids); 0
(Phosphatidylglycerols); EC 2.7.7

(Nucleotidyltransferases); EC 2.7.7.15 (Choline-Phosphate

Cytidylyltransferase); EC 6.- (Fatty Acid Synthetase

Complex)

L75 ANSWER 15 OF 22 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE

ACCESSION NUMBER:

1999:476882 BIOSIS PREV199900476882

DOCUMENT NUMBER: TITLE:

(-)-Hydroxycitric acid does not affect energy expenditure and substrate oxidation in adult males in a post-absorptive

state.

AUTHOR(S):

Kriketos, A. D.; Thompson, H. R.; Greene, H.; Hill, J. O.

CORPORATE SOURCE:

(1)

(1) Center for Human Nutrition, University of Colorado Health Sciences Center, 4200 East Ninth-Avenue, Denver, CO,

80262 USA

SOURCE:

International Journal of Obesity, (Aug., 1999) Vol. 23, No.

8, pp. 867-873. ISSN: 0307-0565.

Jones 09/781491 Page 27

DOCUMENT TYPE: Article LANGUAGE: English SUMMARY LANGUAGE: English

ABSTRACT:

OBJECTIVE: (-)-Hydroxycitric acid ((-)-HCA) is available as a herbal supplement, and promoted as a weight loss agent. It is hypothesized that (-)-HCA can increase fat oxidation by inhibiting citrate lyase, an enzyme which plays a crucial role in energy metabolism during de novo lipogenesis. The indirect inhibition of the cytosolic pool of citrate by (-)-HCA and the subsequent reduction in acetyl coenzyme A and oxaloacetate alters steps in the citric acid cycle that promote fat oxidation. The objective of this study was to determine the effect of (-)-HCA on marker substrates of altered metabolism, as well as on respiratory quotient (RQ) and energy expenditure (EE) in humans, following an overnight fast and during a bout of exercise. HYPOTHESIS OF STUDY: We hypothesized that supplementation with (-)-HCA would result in an increase in fat oxidation and metabolic rate, reflected by an increase in beta-hydroxybutyrate and EE and/or a decrease in RQ. Furthermore, during moderately intense exercise, we hypothesized that (-)-HCA supplementation would increase the rate of lactate conversion to glucose in the liver, with a subsequent reduction of circulating lactate and an elevation of circulating ketone bodies due to the increased partial oxidation of fatty acids (FA) in mitochondria. Studies have examined the fat regulating action of (-)-HCA on steps of the citric acid cycle in rodents showing reductions in body weight and food intake. No studies have investigated the effects of (-)-HCA supplementation in conjunction with a typical daily dietary composition (that is approx 30 - 35% fat) on metabolic processes which could influence body weight regulation in humans. OESIGN This was a double blind, placebo controlled, randomized, crossover study involving three days of (-)-HCA (3.0 g/d) or placebo supplementation. The effects of (-)-HCA supplementation on metabolic parameters with or without moderately intense exercise was studied over four laboratory visits. SUBJECTS: Sedentary adult male subjects (n = 10, age: 22 - 38 y, body mass index (BMI) 22.4 -37.6 kg/m2). MEASUREMENTS: Two of the four visits involved no exercise (Protocol A) with and without (-)-HCA treatment, while the remaining two visits included a moderately intense exercise bout (Protocol B; 30 min at 40% maximal aerobic fitness (VO2max) and 15 min at 60% VO2max) with and without (-)-HCA treatment. EE (by indirect calorimetry) and RQ were measured for 150 min following an overnight fast. Blood samples were collected for the determination of glucose, insulin glucagon, lactate, and beta-hydroxybutyrate concentrations. RESULTS: In a fasted state and following 3 d of (-)-HCA treatment, RQ was not significantly lowered during rest (Protocol A) nor during exercise (Protocol B) compared with the placebo treatment. Treatment with (-)-HCA did not affect EE, either during rest or during moderately intense exercise. Furthermore, the blood substrates measured were not significantly different between treatment groups under the fasting conditions of this study. CONCLUSION: These results do not support the hypothesis that ( )-HCA alters the short-term rate of fat oxidation in the fasting state during rest or moderate exercise, with doses likely to be achieved in humans while subjects maintain a typical Western diet (approx 30 -35% total calories as fat).

CONCEPT CODE: Nutrition - Malnutrition; Obesity \*13203

Biochemical Studies - General \*10060

Respiratory System - General; Methods \*16001

Metabolism - Metabolic Disorders \*13020

BIOSYSTEMATIC CODE: Hominidae 86215 INDEX TERMS: Major Concepts

Nutrition

INDEX TERMS: Parts, Structures, & Systems of Organisms

mitochondria

INDEX TERMS: Chemicals & Biochemicals

acetyl coenzyme A [acetyl coA]; beta-hydroxybutyrate;

citrate lyase; fatty acids; glucagon; glucose; insulin; lactate; levo-hydroxycitric acid: herbal

supplement, weight loss agent

Jones 09/781491 Page 28

INDEX TERMS:

Methods & Equipment

indirect calorimetry: analytical method

INDEX TERMS:

Miscellaneous Descriptors

aerobic fitness; body mass index; energy expenditure; exercise; fasting; fat oxidation; lipogenesis; respiratory

quotient; substrate oxidation

ORGANISM:

Super Taxa

Hominidae: Primates, Mammalia, Vertebrata, Chordata,

Animalia

ORGANISM:

Organism Name

human (Hominidae): adult, male

ORGANISM:

Organism Superterms

Animals; Chordates; Humans; Mammals; Primates; Vertebrates

REGISTRY NUMBER:

27750-10-3 ((-)-HYDROXYCITRIC ACID) 27750-10-3 (LEVO-HYDROXYCITRIC ACID)

9012-83-3 (CITRATE LYASE) 72-89-9 (ACETYL COENZYME A)

72-89-9 (ACETYL COA)

300-85-6 (BETA-HYDROXYBUTYRATE)

113-21-3 (LACTATE) 50-99-7Q (GLUCOSE) 58367-01-4Q (GLUCOSE) 9004-10-8 (INSULIN) 9007-92-5 (GLUCAGON)

L75 ANSWER 16 OF 22 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:252645 BIOSIS

TITLE:

PREV200100252645 Nutritional supplement products containing chromium

picolinate and hydroxycitric acid lead to weight loss in

randomized controlled study.

AUTHOR(S):

Greenberg, Danielle (1); Harris, Rosemarie (1); Komorowski,

James R. (1)

CORPORATE SOURCE:

(1) AMBI Inc., 4 Manhattanville Road, Purchase, NY, 10577

SOURCE:

FASEB Journal (March 7, 2001) Vol. 15, No. 4, pp. A75.

print.

Meeting Info.: Annual Meeting of the Federation of American Societies for Experimental Biology on Experimental Biology

2001 Orlando, Florida, USA March 31-April 04, 2001

ISSN: 0892-6638.

DOCUMENT TYPE:

LANGUAGE:

Conference

English

SUMMARY LANGUAGE: English ABSTRACT:

Chromium picolinate (CrPic) and hydroxycitric acid (HCA) have both been reported to have weight-loss Wenefits. We Mamined the effectiveness of a weight-loss program using nutritional snacks and capsules containing both CrPic and HCA. The USDA Food Guide Pyramid program was used as a control. Subjects (BMI 27 - 40 kg/m2) were assigned to exther a treatment group (n=40) that received dietary supplements in the form of bars, snacks or capsules containing Cr (200 - 400 mcg/day) and HCA (1000 - 2000 mg/day) along with essential vitamins and minerals, or to a control group (n=17) receiving no dietary supplement, for 12 weeks. Both groups followed a dietary program using the USDA Food Guide Pyramid guidelines (1200-1600 kcal/day). Both groups received instructions on following these guidelines, had dietary recall monitored and were recommended exercise by a registered dietitian. Body weight, fasting \*\*\*insulin\*\*\* , cholesterol and blood glucose were measured. Weight consistently and steadily declined in the treatment group with a loss (mean  $4.6,\;$  max 19 1bs) that was significantly greater than in the control group (mean0.8, max 6 lbs; F (1.48) = 4.1, p<0.05). There were no significant changes in fasting insulin, cholesterol or blood glucose in either group. We conclude that CrPic and HCA in combination with other nutrients can be

Jones 09/781491 Page 29

effectively used in a moderate weight loss program under normal living conditions without severe caloric restriction. The use of the combination of these nutrient supplements for weight loss deserves further examination.

CONCEPT CODE: Food Technology - General; Methods \*13502

General Biology - Symposia, Transactions and Proceedings of

Conferences, Congresses, Review Annuals \*00520

Metabolism - General Metabolism; Metabolic Pathways \*13002 Nutrition - General Studies, Nutritional Status and Methods

\*13202

Food Technology - Synthetic, Supplemental and Enrichment

Foods \*13534

BIOSYSTEMATIC CODE: Hominidae 86215

INDEX TERMS: Major Concepts

Foods; Metabolism; Nutrition

INDEX TERMS: Chemicals & Biochemicals

> chromium picolinate: dietary supplement; hydroxycitric acid: dietary supplement; nutritional capsules: dietary

supplement

INDEX TERMS: Miscellaneous Descriptors

caloric restriction; nutritional bar: food supplement;

nutritional snack: food supplement; nutritional

supplements: food supplement; weight loss; Meeting Abstract

ORGANISM: Super Taxa

Hominidae: Primates, Mammalia, Vertebrata, Chordata,

Animalia

ORGANISM: Organism Name

human (Hominidae)

ORGANISM: Organism Superterms

Animals; Chordates; Humans; Mammals; Primates; Vertebrates

REGISTRY NUMBER: 6205-14-7Q (HYDROXYCITRIC ACID)

27750-10-3Q (HYDROXYCITRIC ACID)

L75 ANSWER 17 OF 22 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER:

2000400228 EMBASE

TITLE:

A randomized, double-blind, placebo-controlled trial of a

new weight-reducing agent of natural origin.

AUTHOR: Thom E.

CORPORATE SOURCE: Dr. E. Thom, Parexel Medstat AS, PO Box 210, N-2001

Lillestrom, Norway. erling.thom@parexel.com

SOURCE: Journal of International Medical Research, (2000) 28/5

(229-233).

Refs: 13

ISSN: 0300-0605 CODEN: JIMRBV

COUNTRY: DOCUMENT TYPE: United Kingdom Journal; Article

006 Internal Medicine

FILE SEGMENT:

030 Pharmacology

037 Drug Literature Index 038 Adverse Reactions Titles

039 Pharmacy

LANGUAGE:

English SUMMARY LANGUAGE: English

ABSTRACT:

The efficacy and tolerability of a new weight-reduction agent, based on natural ingredients, was investigated in this randomized, placebo-controlled, double-blind study. The product reduces the absorption of different types of sugar from the gastrointestinal tract. Forty obese volunteers were included in the 12-week study. Body weight, body composition and blood pressure were recorded at baseline and every month during the study. The results show a significant difference in weight reduction in favour of the active group (3.5 kg versus 1.2 kg). Body composition measurements showed that > 85% of the reduction in the active group is fat loss. The tolerability was similar and good in both groups. This product shows promising results and should be studied more extensively at different dose levels.

CONTROLLED TERM:

```
Medical Descriptors:
*weight reduction
*obesity: DT, drug therapy
drug efficacy
drug effect
glucose absorption
stomach absorption
intestine absorption
body weight
body composition
  blood pressure
body fat
drug tolerability
drug mixture
drug formulation
side effect: SI, side effect
human
male
female
clinical article
clinical trial
randomized controlled trial
double blind procedure
controlled study
adult
article
Drug Descriptors:
*natural product: AE, adverse drug reaction
*natural product: CT, clinical trial
*natural product: DT, drug therapy
*natural product: PR, pharmaceutics
*natural product: PD, pharmacology
*natural product: PO, oral drug administration
*suco bloc: AE, adverse drug reaction
*suco bloc: CT, clinical trial
*suco bloc: DT, drug therapy
*suco bloc: PR, pharmaceutics
*suco bloc: PD, pharmacology
*suco bloc: PO, oral drug administration
*antiobesity agent: AE, adverse drug reaction
*antiobesity agent: CT, clinical trial
*antiobesity agent: DT, drug therapy
*antiobesity agent: PR, pharmaceutics
*antiobesity agent: PD, pharmacology
*antiobesity agent: PO, oral drug administration
phaseolus vulgaris extract: AE, adverse drug reaction
phaseolus vulgaris extract: CT, clinical trial
phaseolus vulgaris extract: CB, drug combination
phaseolus vulgaris extract: DT, drug therapy
phaseolus vulgaris extract: PD, pharmacology
phaseolus vulgaris extract: PO, oral drug administration
Garcinia cambogia extract: AE, adverse drug reaction
Garcinia cambogia extract: CT, clinical trial
Garcinia cambogia extract: CB, drug combination
Garcinia cambogia extract: DT, drug therapy
Garcinia cambogia extract: PD, pharmacology
Garcinia cambogia extract: PO, oral drug administration
inulin: AE, adverse drug reaction
inulin: CT, clinical trial
inulin: CB, drug combination
inulin: DT, drug therapy
```

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inulin: PD, pharmacology
                    inulin: PO, oral drug administration
                    hydroxycitric acid: AE, adverse drug reaction
                    hydroxycitric acid: CT, clinical trial
                    hydroxycitric acid: CB, drug combination
                    hydroxycitric acid: DT, drug therapy
                    hydroxycitric acid: PD, pharmacology
                    hydroxycitric acid: PO, oral drug administration
                    amylase inhibitor: PD, pharmacology
                    glycoprotein: AE, adverse drug reaction
                    glycoprotein: CT, clinical trial
                    glycoprotein: CB, drug combination
                    glycoprotein: DT, drug therapy
                    glycoprotein: PD, pharmacology
                    glycoprotein: PO, oral drug administration
                    placebo
                    sugar
                    fat
                    glucose
                    amylase: EC, endogenous compound
                    carbohydrate
                    unclassified drug
                    phaseolamin
                    raftiline
CAS REGISTRY NO.:
                    (inulin) 9005-80-5; (hydroxycitric acid) 27750-10-3
                      6205-14-7; (glucose) 50-99-7, 84778-64-3;
                    (amylase) 9000-90-2, 9000-92-4, 9001-19-8
CHEMICAL NAME:
                    (1) Suco bloc; (2) Phaseolamin; (3) Raftiline
COMPANY NAME:
                    (1) Med Eq (Norway); (2) Leuven Bioproducts (Belgium); (3)
                    Orafti (Belgium)
L75 ANSWER 18 OF 22 EMBASE COPYRIGHT 2002 ELSEVIER SCI
ACCESSION NUMBER:
                    2000298826 EMBASE
                    Current and potential drugs for treatment
AUTHOR:
                    Bray G.A : Greenway F.L.
                    Dr. G.A. Bray, 6400 Perkins Road, Baton Rouge, LX
CORPORATE SOURCE:
                    United States
                    Endocrine Reviews, (1999), 20/6 (805-875)
SOURCE:
                    Refs: 999
                    ISSN: 0163-769X CODEN: ERVIDP
COUNTRY:
                    United States
DOCUMENT TYPE:
                    Journal; General Review
                            Endocrinology
FILE SEGMENT:
                    003
                            Pharmacology
                    030
                    037
                            Drug Literature Index
                    038
                            Adverse Reactions Titles
LANGUAGE:
                    English
CONTROLLED TERM:
                    Medical Descriptors:
                    *obesity: DT, drug therapy
                    *obesity: TH, therapy
                    *drug mechanism
                      pulmonary hypertension: SI, side effect
                      hypertension: SI, side effect
                    gastrointestinal symptom: SI, side effect
                    serotonin syndrome: SI, side effect
                    sleep disorder: SI, side effect
                    anxiety neurosis: SI, side effect
                    behavior modification
                    weight reduction
                    food and drug administration
                    drug marketing
                    cardiotoxicity
```

TITLE:

```
satiety
 energy expenditure
 drug absorption
 food intake
 drug efficacy

    human

 nonhuman
 rat
 review
 priority journal
 Drug Descriptors:
 *antiobesity agent: AE, adverse drug reaction
 *antiobesity agent: DT, drug therapy
 *amylase inhibitor
 *androgen
 *beta adrenergic receptor stimulating agent: PD,
 pharmacology
 *alpha adrenergic receptor stimulating agent: PD,
 pharmacology
 *anorexigenic agent: AE, adverse drug reaction
 *anorexigenic agent: PK, pharmacokinetics
 *anorexigenic agent: PD, pharmacology
 fenfluramine: AE, adverse drug reaction
 fenfluramine: CB, drug combination
 fenfluramine: DT, drug therapy
 fenfluramine: PD, pharmacology
 phentermine: AE, adverse drug reaction
 phentermine: CB, drug combination
 phentermine: DT, drug therapy
 phentermine: PD, pharmacology
 dexfenfluramine: AE, adverse drug reaction
 sibutramine
 mazindol
 tetrahydrolipstatin: AE, adverse drug reaction
 tetrahydrolipstatin: DO, drug dose
 tetrahydrolipstatin: PD, pharmacology
 sucrose polyester: PD, pharmacology
 metformin: PD, pharmacology
 pyruvic acid: PD, pharmacology
 hydroxycitric acid: PD, pharmacology
 chorionic gonadotropin: PD, pharmacology
 prasterone
 testosterone
 thyroid hormone
 ephedrine
 caffeine
 terbutaline: PD, pharmacology
 4 [2 [[2 (3 chlorophenyl) 2 hydroxyethyl]amino]propyl]pheno
 xyacetic acid methyl ester: PD, pharmacology
 4 [2 [(2 hydroxy 2 phenylethyl)amino]propyl]benzoic acid
 methyl ester: PD, pharmacology
 4 [2 [(2 hydroxy 3 phenoxypropyl)amino]ethoxy] n (2
 methoxyethyl) phenoxyacetamide: PD, pharmacology
 4 [3 [bis(beta hydroxyphenethyl)amino]butyl]benzamide: PD,
 pharmacology
 5 [2 [[2 (3 chlorophenyl) 2 hydroxyethyl]amino]propyl] 1,3
 benzodioxole 2,2 dicarboxylic acid: PD, pharmacology
 intermedin
 fluoxetine: PD, pharmacology
 sertraline: PD, pharmacology
 amfepramone
 human growth hormone: PD, pharmacology
 glucose derivative
```

monoamine unindexed drug

4 [2 [(2 hydroxy 2 phenylethyl)amino]propyl]benzoic acid

methyl ester hydrogen maleate

CAS REGISTRY NO.: (fenfluramine) 404-82-0, 458-24-2; (phentermine) 1197-21-3,

122-09-8; (dexfenfluramine) 3239-44-9, 3239-45-0; (sibutramine) 106650-56-0; (mazindol) 22232-71-9;

(tetrahydrolipstatin) 96829-58-2; (metformin) 1115-70-4, 657-24-9; (pyruvic acid) 127-17-3, 19071-34-2, 57-60-3;

(hydroxycitric acid) 27750-10-3,

**6205-14-7**; (chorionic gonadotropin) 9002-61-3;

(prasterone) 53-43-0; (testosterone) 58-22-0; (ephedrine) 299-42-3, 50-98-6; (caffeine) 30388-07-9, 58-08-2; (terbutaline) 23031-25-6; (4 [2 [[2 (3 chlorophenyl) 2 hydroxyethyl]amino]propyl]phenoxyacetic acid methyl ester)

91097-81-3; (4 [2 [(2 hydroxy 2

phenylethyl)amino]propyl]benzoic acid methyl ester)

77955-41-0; (4 [2 [(2 hydroxy 3 phenoxypropyl)amino]ethoxy]

n (2 methoxyethyl)phenoxyacetamide) 129689-30-1; (4 [3 [bis(beta hydroxyphenethyl)amino]butyl]benzamide)

90505-66-1; (5 [2 [[2 (3 chlorophenyl) 2

hydroxyethyl]amino]propyl] 1,3 benzodioxole 2,2

dicarboxylic acid) 138908-40-4; (intermedin) 9002-79-3, 9046-72-4; (fluoxetine) 54910-89-3, 56296-78-7, 59333-67-4; (sertraline) 79617-96-2; (amfepramone) 134-80-5, 90-84-6; (human growth hormone) 12629-01-5; (4 [2 [(2 hydroxy 2 phenylethyl)amino]propyl]benzoic acid methyl ester hydrogen

maleate) 87857-42-9

CHEMICAL NAME: (1) Olean; Brl 35135; Brl 26830a; Ici d7114; Cl 316243; Ro

16 8714

COMPANY NAME: (1) Procter and Gamble (United States)

L75 ANSWER 19 OF 22 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 85245832 EMBASE

DOCUMENT NUMBER:

1985245832

TITLE:

Effect of drugs, peptide hormones and lipogenic precursors

on the relative incorporation of [3H] H2O and carbon into

hepatic cholesterol.

AUTHOR: Bjornsson O.G.; Pullinger C.R.; Gibbons G.F.

CORPORATE SOURCE:

Metabolic Research Laboratory, Nuffield Department of

Clinical Medicine, Radcliffe Infirmary, Oxford OX2 6HE,

United Kingdom

SOURCE: FEBS Letters, (1985) 187/2 (302-306).

CODEN: FEBLAL

COUNTRY:

Netherlands

DOCUMENT TYPE:

Journal

FILE SEGMENT: 037

Drug Literature Index

029 Clinical Biochemistry

048 Gastroenterology

030 Pharmacology

003 Endocrinology

LANGUAGE: English

ABSTRACT:

Measurement of the weight of desmosterol produced during its biosynthesis in the presence of tritiated water and triparanol has permitted a direct determination of the relative flux of caron and tritium (the J/C ratio) into sterol in hepatocytes. The H/C ratio increased with time of incubation irrespective of the nutritional state of the donor animals. This increase was more marked in hepatocytes from starved animals. Pyruvate and lactate increased, and glucagon decreased, the sterol H/C ratio. Addition of pyruvate to incubations containing glucagon resulted in a 32-67% increase in the H/C ratio depending upon nutritional status. Insulin had no effect whilst (-)-hydroxycitrate decreased the ratio by 25%.

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Medical Descriptors: CONTROLLED TERM: \*alpha cyano 3 hydroxycinnamic acid \*cholesterol h 3 \*cholesterol synthesis \*drug accumulation \*drug comparison \*drug identification \*drug interaction \*drug metabolism \*drug tissue level \*starvation diet liver cell priority journal drug analysis nonhuman rat liver animal experiment animal cell in vitro study animal model Drug Descriptors: \*alpha cyano 4 hydroxycinnamic acid \*carbon \*compactin \*desmosterol \*dexamethasone \*glucagon \*hydroxycitric acid \*insulin \*pyruvic acid \*triparanol \*tritium oxide radioisotope (alpha cyano 4 hydroxycinnamic acid) 28166-41-8; (carbon) CAS REGISTRY NO.: 7440-44-0; (compactin) 73573-88-3; (desmosterol) 313-04-2; (dexamethasone) 50-02-2; (glucagon) 11140-85-5, 62340-29-8, 9007-92-5; (hydroxycitric acid) 27750-10-3, 6205-14-7; (insulin) 9004-10-8; (pyruvic acid) 127-17-3, 19071-34-2, 57-60-3; (triparanol) 78-41-1; (tritium oxide) 14940-65-9 COMPANY NAME: Sigma L75 ANSWER 20 OF 22 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V. ACCESSION NUMBER: 83160847 EMBASE 1983160847 DOCUMENT NUMBER: The role of substrate supply in the regulation of TITLE: cholesterol biosynthesis in rat hepatocytes. Pullinger C.R.; Gibbons G.F. AUTHOR: Med. Res. Counc. Lipid Metab. Unit, Hammersmith Hosp., CORPORATE SOURCE: London W12 OHS, United Kingdom Biochemical Journal, (1983) 210/3 (625-632). SOURCE: CODEN: BIJOAK United Kingdom COUNTRY: DOCUMENT TYPE: Journal FILE SEGMENT: 029 Clinical Biochemistry LANGUAGE: English

Compactin, (-)-hydroxycitrate and dexamethasone gave rise to a decrease in the rate of cholesterol production in hepatocytes from fed rats by interfering with the flow of substrate into the sterol biosynthetic pathway. The cells responded

ABSTRACT:

to the deficit of biosynthetic sterol by increasing the activity of hydroxymethylglutaryl-CoA reductase (HMG-CoA reductase). Compactin and (-)-hydroxycitrate gave similar results in hepatocytes from rats starved for 24 h but in this case dexamethasone had no significant effect. Exogenous oleate interferes with the production of carbohydrate-derived acetyl-CoA and also gives rise initially to opposing effects on the rate of sterol synthesis and HMG-CoA reductase activity. Over a longer period, however, oleate itself was capable of replacing carbohydrate as the major source of carbon for sterol synthesis. The increase in HMG-CoA reductase activity observed when liver cells were incubated in the presence of compactin, (-)-hydroxycitrate or oleate could be partially reversed by the simultaneous presence of glucagon. Under some physiological conditions, a deficiency of biosynthetic cholesterol or of a related precursor may lead to an increase in the activity of HMG-CoA reductase.

CONTROLLED TERM: Medical Descriptors:

liver cell animal cell nonhuman rat liver

Drug Descriptors:
\*cholesterol
\*compactin

\*dexamethasone
\*hydroxycitric acid

glucagon

hydroxymethylglutaryl coenzyme a reductase

oleic acid

CAS REGISTRY NO.:

(cholesterol) 57-88-5; (compactin) 73573-88-3; (dexamethasone) 50-02-2; (hydroxycitric acid)

**27750-10-3**, **6205-14-7**; (glucagon)

11140-85-5, 62340-29-8, 9007-92-5; (hydroxymethylglutaryl coenzyme a reductase) 37250-24-1; (oleic acid) 112-80-1,

115-06-0

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=> s 27750-10-3 or 6205-14-7

1 27750-10-3

(27750-10-3/RN) 1 6205-14-7 (6205-14-7/RN) 2 27750-10-3 OR 6205-14-7

L76

=> d ide 176 1-2; fil hom

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RN **27750-10-3** REGISTRY

CN D-erythro-Pentaric acid, 3-C-carboxy-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES: CN (-)-2-Hydroxyc

CN (-)-2-Hydroxycitric acid CN (-)-Hydroxycitric acid

CN Citric acid, 2-hydroxy-, (-)-

CN Garcinia acid

CN Hydroxycitric acid

FS STEREOSEARCH

DR 4373-35-7

MF C6 H8 O8

CI COM

LC STN Files: ADISNEWS, AGRICOLA, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CHEMCATS, CIN, DDFU, DRUGU, EMBASE, HODOC\*, IPA, NAPRALERT, NIOSHTIC, PROMT, TOXCENTER, USPATFULL (\*File contains numerically searchable property data)

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124 REFERENCES IN FILE CAPLUS (1962 TO DATE)

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RN **6205-14-7** REGISTRY

CN Pentaric acid, 3-C-carboxy-2-deoxy- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 1,2,3-Propanetricarboxylic acid, 1,2-dihydroxy- (7CI, 8CI)

OTHER NAMES:

CN Citric acid, hydroxy-

CN Hydroxycitric acid

FS 3D CONCORD

MF C6 H8 O8

CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CIN, CSCHEM, CSNB, EMBASE, MEDLINE, PROMT, TOXCENTER, USPATFULL

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